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chain nodes :
11  12  13  14  15
ring nodes :
1  2  3  4  5  6  7  8  9  10
chain bonds :
1-13  2-15  6-11  9-12  10-14
ring bonds :
1-2  1-6  2-3  3-4  4-5  4-7  5-6  5-10  7-8  8-9  9-10
exact/norm bonds :
1-2  1-6  1-13  2-3  2-15  3-4  4-5  4-7  5-6  5-10  6-11  7-8  8-9  9-10  9-12
10-14
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G1:H,Ak

Match level :

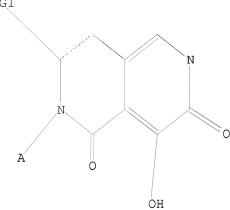
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS

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L1 HAS NO ANSWERS

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12 SEA SSS SAM L1

=> s 11 full

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100.0% PROCESSED 12511 ITERATIONS 245 ANSWERS

SEARCH TIME: 00.00.01

245 SEA SSS FUL L1 L3

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ACCESSION NUMBER: 2006:1205598 CAPLUS

DOCUMENT NUMBER: 145:505478

TITLE: Preparation of tricyclic naphthyridine derivatives, as

HIV integrase inhibitors

INVENTOR(S): Vacca, Joseph P.; Wai, John S.; Payne, Linda S.;

Isaacs, Richard C. A.; Han, Wei; Egbertson, Melissa;

Pracitto, Richard

PATENT ASSIGNEE(S): Merck & Co., Inc., USA SOURCE: PCT Int. Appl., 122pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

GΙ

PATE	KIND DATE				APPLICATION NO.						DATE							
_	2006121831 2006121831								1	wo 2	 7-00c		20060505					
		CN, GE,	CO, GH,	CR, GM,	CU, HR,	CZ, HU,	AU, DE, ID, LT,	DK, IL,	DM, IN,	DZ, IS,	EC, JP,	EE, KE,	EG, KG,	ES, KM,	FI, KN,	GB, KP,	GD, KR,	
		SG,	SK,	SL,		SY,	ΝΖ, ΤJ,	•		•			•	•	•		•	
		IS, CF,	IT, CG,	LT,	LU, CM,	LV, GA,	CZ, MC, GN, NA,	NL, GQ,	PL, GW,	PT, ML,	RO, MR,	SE, NE,	SI, SN,	SK, TD,	TR, TG,	BF, BW,	BJ, GH,	
	AU 2006244424									EA, EP, OA AU 2006-244424 CA 2006-2607151								
EP 1	R:	AT, IS,	BE, IT,	BG, LI,	CH, LT,	CY,	20080 CZ, LV,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,	
	BA, HR, MK, RIORITY APPLN. INFO.:						145:	5054	US 2005-679431P WO 2006-US17369									

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Tricyclic compds. I [R1 = (un)substituted alkyl; R2 = H, alkyl; R3 = H, halo, CN, halo/alkyl, SO2NH2 and derivs., NH2 and derivs., etc.; when R2 and R3 are attached to a double bond, then R2C:CR3 = (un)substituted Ph, 5- to 6-membered heteroaryl; A = (un)substituted 5- to 9-membered saturated or mono-unsatd. heterocyclyl containing 2-4 heteroatoms selected from N, O, S, in which each S is optionally oxidized to SO or SO2; ; and their pharmaceutically acceptable salts] were prepared as HIV integrase and HIV replication inhibitors. For example, diazocinonaphthyridine II was prepared by coupling acid III (preparation given) with N-[2-(benzyloxy)ethyl]-4-[(tert-butyldimethylsilyl)oxy]butan-1-amine (preparation given), removal of tert-butyldimethylsilyl group, treatment of the alc. with methanesulfonic anhydride, cyclization in the presence of CsCO3, and deprotection in the presence of HBr in AcOH (no data for the first two steps). I displayed IC50's <1 μM in an HIV integrase assay. Pyrazinopyrimidines I

exhibited IC95's of <10 μM for the inhibition of acute HIV infection of T-lymphoid cells. I exhibited no cytotoxicity at concns. $\leq \! 10$ μM . I are useful in the prevention and treatment of infection by HIV and in the prevention, delay in the onset, and treatment of AIDS. I can be employed in pharmaceutical compns., optionally in combination with other antivirals, immunomodulators, antibiotics or vaccines.

865300-53-4P, Methyl 6-(3-chloro-4-fluorobenzyl)-3,4-dihydroxy-5-oxo-5,6,7,8-tetrahydro-2,6-naphthyridine-1-carboxylate
915065-29-1P, Methyl 2-(4-chlorobutyl)-6-(3-chloro-4-fluorobenzyl)4-hydroxy-3,5-dioxo-2,3,5,6,7,8-hexahydro-2,6-naphthyridine-1-carboxylate
915065-30-4P, Methyl 6-(3-chloro-4-fluorobenzyl)-4-hydroxy-2-[4-(methylamino)butyl]-3,5-dioxo-2,3,5,6,7,8-hexahydro-2,6-naphthyridine-1-carboxylate 915065-31-5P, 6-(3-chloro-4-fluorobenzyl)-4-hydroxy-2-[4-(methylamino)butyl]-3,5-dioxo-2,3,5,6,7,8-hexahydro-2,6-naphthyridine-1-carboxylic acid hydrochloride 915065-35-9P,

10-(3-Chloro-4-fluorobenzyl)-8-hydroxy-2-(4-methoxybenzyl)-2,3,4,5,11,12-hexahydro-[1,4]diazepino[2,1-a][2,6]naphthyridine-1,7,9(10H)-trione 915065-36-0P, Methyl 2-(2-tert-butoxy-2-oxoethyl)-6-(3-chloro-4-fluorobenzyl)-4-hydroxy-3,5-dioxo-2,3,5,6,7,8-hexahydro-2,6-naphthyridine-1-carboxylate 915065-66-6P, Methyl 2-(allyl)-6-(3-chloro-4-fluorobenzyl)-4-hydroxy-3,5-dioxo-2,3,5,6,7,8-hexahydro-2,6-naphthyridine-1-carboxylate

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(intermediate; preparation of tricyclic naphthyridines as HIV integrase inhibitors)

RN 865300-53-4 CAPLUS

ΙT

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 915065-29-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 2-(4-chlorobutyl)-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-[4-(methylamino)butyl]-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 915065-31-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-[4-(methylamino)butyl]-3,5-dioxo-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 915065-35-9 CAPLUS

CN [1,4]Diazepino[2,1-a][2,6]naphthyridine-1,7,9(10H)-trione, 10-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,5,11,12-hexahydro-8-hydroxy-2-[(4-methoxyphenyl)methyl]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ \text{MeO} & & & \\ & & & \\ \text{CH}_2 & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

RN 915065-36-0 CAPLUS

CN 2,6-Naphthyridine-2(3H)-acetic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-5,6,7,8-tetrahydro-4-hydroxy-1-(methoxycarbonyl)-3,5-dioxo-, 1,1-dimethylethyl ester (CA INDEX NAME)

RN 915065-66-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-2-(2-propen-1-yl)-, methyl ester (CA INDEX NAME)

IT 915065-12-2P, (6R)-11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-6 (hydroxymethyl)-2-methyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]2,6-naphthyridine-1,8,10(11H)-trione 915065-27-9P,
9-(3-Chloro-4-fluorobenzyl)-7-hydroxy-3-(acetyloxymethyl)-3,4,10,11tetrahydro-[1,4]oxazino[3,4-a][2,6]naphthyridine-1,6,8(9H)-trione
RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological
activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or
reagent); USES (Uses)

(preparation of tricyclic naphthyridines as HIV integrase inhibitors)

RN 915065-12-2 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-(hydroxymethyl)-2-methyl-, (6R)- (CA INDEX NAME)

Absolute stereochemistry.

RN 915065-27-9 CAPLUS

CN [1,4]Oxazino[3,4-a][2,6]naphthyridine-1,6,8(9H)-trione,

3-[(acetyloxy)methyl]-9-[(3-chloro-4-fluorophenyl)methyl]-3,4,10,11-tetrahydro-7-hydroxy- (CA INDEX NAME)

ΙT 915064-71-0P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-methyl-3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-naphthyridine-1,8,10(11H)-trione 915064-72-1P, 11-(4-Fluorobenzyl)-9-hydroxy-2methyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione 915064-73-2P, 10-(3-Chloro-4-fluorobenzyl)-8hydroxy-2-methyl-2, 3, 4, 5, 11, 12-hexahydro-[1, 4]diazepino[2, 1a][2,6]naphthyridine-1,7,9(10H)-trione 915064-74-3P, 10-(3-Chloro-4-fluorobenzyl)-8-hydroxy-2,3,4,5,11,12-hexahydro-[1,4] diazepino [2,1-a] [2,6] naphthyridine-1,7,9(10H)-trione 915064-75-4P, 9-(3-Chloro-4-fluorobenzyl)-7-hydroxy-3,4,10,11tetrahydro-2H-pyrazino[2,1-a][2,6]naphthyridine-1,6,8(9H)-trione 915064-76-5P, 9-(3-Chloro-4-fluorobenzyl)-7-hydroxy-2-methyl-3,4,10,11-tetrahydro-2H-pyrazino[2,1-a][2,6]naphthyridine-1,6,8(9H)-trione 915064-77-6P, 9-(3-Chloro-4-fluorobenzyl)-7-hydroxy-3,4,10,11tetrahydro-[1,4]oxazino[3,4-a][2,6]naphthyridine-1,6,8(9H)-trione 915064-78-7P, 8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1,3-dimethyl-1,8,9,10-tetrahydroimidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione 915064-79-8P, 8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1,3,3trimethyl-1,8,9,10-tetrahydroimidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)trione 915064-80-1P, 11-(3-Chloro-4-fluorobenzy1)-9-hydroxy-2-[2-(acetyloxy) ethyl]-3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6naphthyridine-1,8,10(11H)-trione 915064-81-2P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-(2-hydroxyethyl)-3,4,5,6,12,13hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione 915064-82-3P 915064-83-4P 915064-84-5P 915064-85-6P 915064-86-7P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(morpholin-4-yl)ethyl]-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione 915064-87-8P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(pyrrolidin-1-y1)ethy1]-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione 915064-88-9P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(piperidin-1-yl)ethyl]-3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-naphthyridine-1,8,10(11H)-trione 915064-89-0P, 11-(3-Chloro-4-fluorobenzyl)-9hydroxy-2-(2-aminoethy1)-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione 915064-90-3P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(acetylamino)ethyl]-3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-naphthyridine-1,8,10(11H)-trione 915064-91-4P, 11-(3-Chloro-4-fluorobenzyl)-9hydroxy-2-[2-[(methylsulfonyl)amino]ethyl]-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione 915064-92-5P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(methoxy) ethyl] -3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-hexahydro-2H-[1, 4]-2, 6-hexnaphthyridine-1,8,10(11H)-trione 915064-93-6P 915064-94-7P 915064-95-8P, 11-(3-Chloro-4-fluorobenzyl)-

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9-hydroxy-2-methyl-4-(pyrrolidin-1-yl)-3, 4, 5, 6, 12, 13-hexahydro-2H-
[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione
915064-96-9P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-methyl-4-
(morpholin-4-yl)-3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-hexahydro-2H-[1, 4]-2, 6-hexahydro-2H-[1, 4]-2
naphthyridine-1, 8, 10(11H)-trione 915064-97-0P,
11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(morpholin-4-yl)ethyl]-6-
methyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-
1,8,10(11H)-trione 915064-98-1P, 11-(3-Chloro-4-fluorobenzyl)-9-
hydroxy-2-[2-(piperidin-1-y1)ethyl]-6-methyl-3, 4, 5, 6, 12, 13-hexahydro-2H-
[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione
915064-99-2P, 11-(3-Chloro-4-fluorobenzy1)-9-hydroxy-2-[2-
(pyrrolidin-1-y1)ethy1]-6-methy1-3,4,5,6,12,13-hexahydro-2H-
[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione
915065-00-8P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(1H-
pyrazol-1-yl)ethyl]-6-methyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-
a]-2,6-naphthyridine-1,8,10(11H)-trione 915065-01-9P,
11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2-[2-(1H-imidazol-1-yl)ethyl]-6-
methyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-
1,8,10(11H)-trione 915065-02-0P, 11-(3-Chloro-4-fluorobenzy1)-
5,9-dihydroxy-2-methyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-
2,6-naphthyridine-1,8,10(11H)-trione 915065-03-1P,
(4R)-11-(3-Chloro-4-fluorobenzyl)-4,9-dihydroxy-2,5,5-trimethyl-
3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-naphthyridine-
1,8,10(11H)-trione 915065-04-2P, (4S)-11-(3-Chloro-4-
fluorobenzy1)-4,9-dihydroxy-2,5,5-trimethyl-3,4,5,6,12,13-hexahydro-2H-
[1,4] diazocino [2,1-a]-2, 6-naphthyridine-1, 8, 10 (1H) -trione
915065-05-3P 915065-07-5P, 11-(4-Fluorobenzyl)-4,9-
dihydroxy-2,5,5-trimethyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-
2,6-naphthyridine-1,8,10(11H)-trione 915065-08-6P,
(4R,6S)-11-(3-Chloro-4-fluorobenzyl)-4,9-dihydroxy-2,6-dimethyl-
3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-
1,8,10(11H)-trione 915065-09-7P, (4S,6S)-11-(3-Chloro-4-
fluorobenzyl)-4,9-dihydroxy-2,6-dimethyl-3,4,5,6,12,13-hexahydro-2H-
[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione
915065-10-0P, 11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2,4,4-
trimethyl-3,4,5,6,12,13-hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-
naphthyridine-1,8,10(11H)-trione 915065-11-1P,
11-(3-Chloro-4-fluorobenzyl)-9-hydroxy-2,4-dimethyl-3,4,5,6,12,13-
hexahydro-2H-[1,4]diazocino[2,1-a]-2,6-naphthyridine-1,8,10(11H)-trione
915065-13-3P, (6R)-11-(4-Fluorobenzyl)-9-hydroxy-6-(hydroxymethyl)-
2-methyl-3, 4, 5, 6, 12, 13-hexahydro-2H-[1, 4]diazocino[2, 1-a]-2, 6-
naphthyridine-1, 8, 10(11H)-trione 915065-14-4P
915065-15-5P 915065-16-6P 915065-17-7P
915065-18-8P, 8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1-(2-
hydroxyethyl)-3,3-dimethyl-1,8,9,10-tetrahydroimidazo[2,1-
a][2,6]naphthyridine-2,5,7(3H)-trione 915065-19-9P,
8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1-(3-hydroxypropyl)-3,3-dimethyl-
1,8,9,10-tetrahydroimidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione
915065-20-2P, 8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1-(4-
hydroxybutyl)-3,3-dimethyl-1,8,9,10-tetrahydroimidazo[2,1-
a][2,6]naphthyridine-2,5,7(3H)-trione 915065-21-3P,
8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1-(2-acetyloxyethyl)-3,3-dimethyl-
1,8,9,10-tetrahydroimidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione
915065-22-4P, 8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1-(3-
acetyloxypropyl)-3,3-dimethyl-1,8,9,10-tetrahydroimidazo[2,1-a]-2,6-
naphthyridine-2,5,7(3H)-trione 915065-23-5P,
8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1-(4-acetyloxybutyl)-3,3-dimethyl-
1,8,9,10-tetrahydroimidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione
915065-24-6P, 8-(3-Chloro-4-fluorobenzyl)-6-hydroxy-1,3-bis(2-
\label{localization} \verb|hydroxyethy1| -3 - methy1 -1, 8, 9, 10 - tetrahydroimidazo[2, 1-a][2, 6] | naphthyridine-localization and the statement of the stateme
2,5,7(3H)-trione 915065-25-7P, 11-(3-Chloro-4-fluorobenzyl)-9-
hydroxy-3,4,5,6,12,13-hexahydro-[1,4]oxazocino[3,4-a]-2,6-naphthyridine-
```

1,8,10(11H)-trione 915065-26-8P, 10-(3-Chloro-4-fluorobenzyl)-8hydroxy-4,5,11,12-tetrahydro-3H-[1,4]oxazepino[3,4-a][2,6]naphthyridine-1,7,9(10H)-trione 915065-28-0P, 9-(3-Chloro-4-fluorobenzyl)-7hydroxy-3-(hydroxymethyl)-3,4,10,11-tetrahydro-[1,4]oxazino[3,4a][2,6]naphthyridine-1,6,8(9H)-trione 915095-68-0P 915095-69-1P 915095-70-4P 915095-71-5P 915095-72-6P 915095-73-7P RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of tricyclic naphthyridines as HIV integrase inhibitors) RN 915064-71-0 CAPLUS CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2methyl- (CA INDEX NAME)

RN 915064-72-1 CAPLUS
CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione,
11-[(4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-methyl(CA INDEX NAME)

RN 915064-73-2 CAPLUS
CN [1,4]Diazepino[2,1-a][2,6]naphthyridine-1,7,9(10H)-trione,
10-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,5,11,12-hexahydro-8-hydroxy-2methyl- (CA INDEX NAME)

RN 915064-74-3 CAPLUS

CN [1,4]Diazepino[2,1-a][2,6]naphthyridine-1,7,9(10H)-trione, 10-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,5,11,12-hexahydro-8-hydroxy-(CA INDEX NAME)

RN 915064-75-4 CAPLUS

CN 2H-Pyrazino[2,1-a][2,6]naphthyridine-1,6,8(9H)-trione, 9-[(3-chloro-4-fluorophenyl)methyl]-3,4,10,11-tetrahydro-7-hydroxy- (CA INDEX NAME)

RN 915064-76-5 CAPLUS

CN 2H-Pyrazino[2,1-a][2,6]naphthyridine-1,6,8(9H)-trione, 9-[(3-chloro-4-fluorophenyl)methyl]-3,4,10,11-tetrahydro-7-hydroxy-2-methyl- (CA INDEX NAME)

RN 915064-77-6 CAPLUS

CN [1,4]Oxazino[3,4-a][2,6]naphthyridine-1,6,8(9H)-trione, 9-[(3-chloro-4-fluorophenyl)methyl]-3,4,10,11-tetrahydro-7-hydroxy- (CA INDEX NAME)

RN 915064-78-7 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-1,3-dimethyl-(CA INDEX NAME)

RN 915064-79-8 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-1,3,3-trimethyl-(CA INDEX NAME)

RN 915064-80-1 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 2-[2-(acetyloxy)ethyl]-11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy- (CA INDEX NAME)

RN 915064-81-2 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-(2-hydroxyethyl)- (CA INDEX NAME)

RN 915064-82-3 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-2-acetic acid, 11-[(3-chloro-4-fluorophenyl)methyl]-1,3,4,5,6,8,10,11,12,13-decahydro-9-hydroxy-1,8,10-trioxo- (CA INDEX NAME)

RN 915064-83-4 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-2-acetamide, 11-[(3-chloro-4-fluorophenyl)methyl]-1,3,4,5,6,8,10,11,12,13-decahydro-9-hydroxy-N,N-dimethyl-1,8,10-trioxo- (CA INDEX NAME)

$$\begin{array}{c} C1 \\ Me_2N-C-CH_2 \\ N \\ O \\ O \\ O \end{array}$$

RN 915064-84-5 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-2-acetamide, 11-[(3-chloro-4-fluorophenyl)methyl]-1,3,4,5,6,8,10,11,12,13-decahydro-9-hydroxy-N-methyl-1,8,10-trioxo- (CA INDEX NAME)

RN 915064-85-6 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-2-acetamide, 11-[(3-chloro-4-fluorophenyl)methyl]-1,3,4,5,6,8,10,11,12,13-decahydro-9-hydroxy-1,8,10-trioxo- (CA INDEX NAME)

RN 915064-86-7 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-[2-(4-morpholinyl)ethyl]- (CA INDEX NAME)

RN 915064-87-8 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-[2-(1-pyrrolidinyl)ethyl]- (CA INDEX NAME)

RN 915064-88-9 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-[2-(1-piperidinyl)ethyl]- (CA INDEX NAME)

RN 915064-89-0 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 2-(2-aminoethyl)-11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy- (CA INDEX NAME)

$$H_2N-CH_2-CH_2$$
 N
 O
 O
 O
 O
 O

RN 915064-90-3 CAPLUS

CN Acetamide, N-[2-[11-[(3-chloro-4-fluorophenyl)methyl]-1,3,4,5,6,8,10,11,12,13-decahydro-9-hydroxy-1,8,10-trioxo-2H-[1,4]diazocino[2,1-a][2,6]naphthyridin-2-yl]ethyl]- (CA INDEX NAME)

RN 915064-91-4 CAPLUS

CN Methanesulfonamide, N-[2-[11-[(3-chloro-4-fluorophenyl)methyl]-1,3,4,5,6,8,10,11,12,13-decahydro-9-hydroxy-1,8,10-trioxo-2H-[1,4]diazocino[2,1-a][2,6]naphthyridin-2-yl]ethyl]- (CA INDEX NAME)

RN 915064-92-5 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-(2-methoxyethyl)- (CA INDEX NAME)

$$\begin{array}{c} \text{C1} \\ \text{MeO-CH}_2\text{-CH}_2 \\ \text{O} \\ \text{N} \\ \text{OH} \end{array}$$

RN 915064-93-6 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2-methyl- (CA INDEX NAME)

RN 915064-94-7 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,4,8,10(3H,11H)-tetrone, 11-[(3-chloro-4-fluorophenyl)methyl]-5,6,12,13-tetrahydro-9-hydroxy-2-methyl- (CA INDEX NAME)

RN 915064-95-8 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-methyl-4-(1-pyrrolidinyl)- (CA INDEX NAME)

RN 915064-96-9 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-methyl-4-(4-morpholinyl)- (CA INDEX NAME)

RN 915064-97-0 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-methyl-2-[2-(4-morpholinyl)ethyl]- (CA INDEX NAME)

RN 915064-98-1 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-methyl-2-[2-(1-piperidinyl)ethyl]- (CA INDEX NAME)

RN 915064-99-2 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-methyl-2-[2-(1-pyrrolidinyl)ethyl]- (CA INDEX NAME)

RN 915065-00-8 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-methyl-2-[2-(1H-pyrazol-1-yl)ethyl]- (CA INDEX NAME)

RN 915065-01-9 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-[2-(1H-imidazol-1-yl)ethyl]-6-methyl- (CA INDEX NAME)

RN 915065-02-0 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-5,9-dihydroxy-2-methyl- (CA INDEX NAME)

RN 915065-03-1 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2,5,5-trimethyl-, (4R)- (CA INDEX NAME)

Absolute stereochemistry.

RN 915065-04-2 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2,5,5-trimethyl-, (4S)- (CA INDEX NAME)

Absolute stereochemistry.

RN 915065-05-3 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2,5,5-trimethyl- (CA INDEX NAME)

RN 915065-07-5 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2,5,5-trimethyl- (CA INDEX NAME)

RN 915065-08-6 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2,6-dimethyl-, (4R,6S)- (CA INDEX NAME)

Absolute stereochemistry.

RN 915065-09-7 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-4,9-dihydroxy-2,6-dimethyl-, (4S,6S)- (CA INDEX NAME)

Absolute stereochemistry.

RN 915065-10-0 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2,4,4-trimethyl- (CA INDEX NAME)

RN 915065-11-1 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2,4-dimethyl- (CA INDEX NAME)

RN 915065-13-3 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-(hydroxymethyl)-2-methyl-, (6R)- (CA INDEX NAME)

Absolute stereochemistry.

RN 915065-14-4 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-methyl-6-methylene- (CA INDEX NAME)

RN 915065-15-5 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-1(5H)-acetamide, 8-[(3-chloro-4-fluorophenyl)methyl]-2,3,7,8,9,10-hexahydro-6-hydroxy-N,N,3,3-tetramethyl-2,5,7-trioxo- (CA INDEX NAME)

RN 915065-16-6 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-1(5H)-acetamide, 8-[(3-chloro-4-fluorophenyl)methyl]-2,3,7,8,9,10-hexahydro-6-hydroxy-N,3,3-trimethyl-2,5,7-trioxo-(CA INDEX NAME)

RN 915065-17-7 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-1(5H)-acetamide, 8-[(3-chloro-4-fluorophenyl)methyl]-2,3,7,8,9,10-hexahydro-6-hydroxy-3,3-dimethyl-2,5,7-trioxo- (CA INDEX NAME)

$$\begin{array}{c|c} C1 \\ H_2N-C-CH_2 \\ \hline \\ N \\ \hline \\ Me \\ O \end{array}$$

RN 915065-18-8 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-1-(2-hydroxyethyl)-3,3-dimethyl- (CA INDEX NAME)

RN 915065-19-9 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-1-(3-hydroxypropyl)-3,3-dimethyl- (CA INDEX NAME)

RN 915065-20-2 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-1-(4-hydroxybutyl)-3,3-dimethyl- (CA INDEX NAME)

RN 915065-21-3 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 1-[2-(acetyloxy)ethyl]-8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-3,3-dimethyl- (CA INDEX NAME)

RN 915065-22-4 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 1-[3-(acetyloxy)propyl]-8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-3,3-dimethyl- (CA INDEX NAME)

RN 915065-23-5 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 1-[4-(acetyloxy)butyl]-8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-3,3-dimethyl- (CA INDEX NAME)

RN 915065-24-6 CAPLUS

CN Imidazo[2,1-a][2,6]naphthyridine-2,5,7(3H)-trione, 8-[(3-chloro-4-fluorophenyl)methyl]-1,8,9,10-tetrahydro-6-hydroxy-1,3-bis(2-hydroxyethyl)-3-methyl- (CA INDEX NAME)

RN 915065-25-7 CAPLUS

CN [1,4]Oxazocino[3,4-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-(CA INDEX NAME)

RN 915065-26-8 CAPLUS

CN 3H-[1,4]Oxazepino[3,4-a][2,6]naphthyridine-1,7,9(10H)-trione, 10-[(3-chloro-4-fluorophenyl)methyl]-4,5,11,12-tetrahydro-8-hydroxy- (CA INDEX NAME)

RN 915065-28-0 CAPLUS

CN [1,4]Oxazino[3,4-a][2,6]naphthyridine-1,6,8(9H)-trione, 9-[(3-chloro-4-fluorophenyl)methyl]-3,4,10,11-tetrahydro-7-hydroxy-3-(hydroxymethyl)- (CA INDEX NAME)

RN 915095-68-0 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-(hydroxymethyl)-2-methyl-, (6S,7S)- (9CI) (CA INDEX NAME)

RN 915095-69-1 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2,6-dimethyl-, (6R,7R)-rel- (9CI) (CA INDEX NAME)

RN 915095-70-4 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2,6-dimethyl-, (6R,7S)-rel- (9CI) (CA INDEX NAME)

RN 915095-71-5 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-(2-hydroxyethyl)-6-methyl-, (6R,7R)-rel- (9CI) (CA INDEX NAME)

RN 915095-72-6 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-2-(2-hydroxyethyl)-6-methyl-, (6R,7S)-rel- (9CI) (CA INDEX NAME)

RN 915095-73-7 CAPLUS

CN 2H-[1,4]Diazocino[2,1-a][2,6]naphthyridine-1,8,10(11H)-trione, 11-[(3-chloro-4-fluorophenyl)methyl]-3,4,5,6,12,13-hexahydro-9-hydroxy-6-(hydroxymethyl)-2-methyl-, (6S,7R)- (9CI) (CA INDEX NAME)

ANSWER 2 OF 5 CAPLUS COPYRIGHT 2008 ACS on STN L4 2006:1066462 CAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 145:419116 Preparation of crystalline 6-(3-chloro-4-fluorobenzyl)-TITLE: 4-hydroxy-2-isopropyl-N, N-dimethyl-3, 5-dioxo-2,3,5,6,7,8-hexahydro-2,6-naphthyridine-1-carboxamide sodium salt as an HIV integrase inhibitor. INVENTOR(S): Matty, Louis, Jr.; Wang, Yaling PATENT ASSIGNEE(S): Merck & Co., Inc., USA PCT Int. Appl., 23pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE ----_____ _____ WO 2006107478 A2 WO 2006-US7716 20060303 20061012 WO 2006107478 A3 20070712 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AP, EA, EP, OA US 2005-659952P P 20050309 PRIORITY APPLN. INFO.: Title compound (I) having a specified X-ray powder diffraction pattern was prepared as an HIV integrase inhibitor useful for the prophylaxis or treatment of HIV infection or for the prophylaxis, treatment or delay in the onset of AIDS (no data). I was prepared from Et 6-(3-chloro-4fluorobenzyl)-4-hydroxy-5-oxo-5,6,7,8-tetrahydro-2,6-naphthyridine-1carboxylate in 7 steps. ΙT 865300-19-2P RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of crystalline (chlorofluorobenzyl) hydroxyisopropyldioxohexahydronap hthyridinedimethylamide sodium salt as an HIV integrase inhibitor) 865300-19-2 CAPLUS RN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-CN

2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo-,

sodium salt (1:1) (CA INDEX NAME)

Na

IT 865298-83-5P 865300-62-5P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of crystalline

(chlorofluorobenzyl) hydroxyisopropyldioxohexahydronap

hthyridinedimethylamide sodium salt as an HIV integrase inhibitor)

RN 865298-83-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo-(CA INDEX NAME)

RN 865300-62-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

L4 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2005:1026950 CAPLUS

DOCUMENT NUMBER: 143:326344

TITLE: Preparation of hydroxynaphthyridinediones as HIV

integrase inhibitors

INVENTOR(S): Han, Wei; Egbertson, Melissa; Wai, John S.; Zhuang,

Linghang; Ruzek, Rowena D.; Perlow, Debra S.; Isaacs,

Richard C. A.; Cameron, Mark; Foster, Bruce S.;

Dolling, Ulf H.; Hoerrner, R. Scott; Obligado, Vanessa E.; Neilson, Lou Anne; Kim, Boyoung; Payne, Linda S.;

Morrisette, Matthew M.; Williams, Peter D.; Pye,

Philip J.; Angelaud, Remy; Mancheno, Danny E.; Askin,

David

PATENT ASSIGNEE(S): Merck & Co., Inc., USA PCT Int. Appl., 193 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.						KIND DATE		APPLICATION NO.					DATE					
WO	WO 2005087768					A1 20050922			WO 2005-US7772					20050309					
	W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,		
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
		GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	KΖ,	LC,		
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,		
		•	•			•	PL,		•				•	•	•	•	•		
		SY,	ΤJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW	
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MΖ,	NΑ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,		
		ΑZ,	BY,	KG,	KΖ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,		
		EE,	ES,	FΙ,	FR,	GB,	GR,	ΗU,	ΙE,	IS,	ΙΤ,	LT,	LU,	MC,	NL,	PL,	PT,		
		RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,		
		,	,	- ,	TD,	_													
	AU 2005222391					20050922													
CA	2558026				A1		2005	0922		CA 2005-2558026									
EP								EP 2005-725120											
	R:						CZ,												
		IS,	ΙΤ,	LI,	LT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	AL,	BA,		
		•	LV,	MK,	YU A														
	CN 1930160						2007						7404		_				
	JP 2007528401																		
	US 20070179196																		
	IN 2006DN04787					A 20070831													
RIORIT	IORITY APPLN. INFO.:									US 2004-551625P									
													34P			0041			
	HED COLLDON (C)									WO 2005-US7772					W 20050309				

OTHER SOURCE(S): MARPAT 143:326344

GΙ

$$X^{1}$$
 X^{2}
 X^{3}
 X^{2}
 X^{3}
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 X^{4}
 X^{5}
 X^{7}
 X^{7

The title compds. I [X1, X2 = H, alkyl, OH, etc.; or X1 and X2 can AΒ together form methylenedioxy or ethylenedioxy (if located on adjacent carbons in the Ph ring); X3 = H, alkyl, alkoxy, haloalkyl, haloalkoxy, halo; R4 = alkyl, (un)substituted cycloalkyl, heteroaryl, etc.; R5 = H, alkyl, cycloalkyl, etc.] which are inhibitors of HIV integrase and inhibitors of HIV replication, and therefore are useful in the prevention and treatment of infection by HIV and in the prevention, delay in the onset, and treatment of AIDS, were prepared E.g., a multi-step synthesis of II, starting from piperidin-2-one and 4-fluorobenzyl bromide, was given. The representative compds. I (including II) were tested in the integrase assay and were found to have IC50's less than about 1 μM . The compds. I can be employed against HIV infection and AIDS as compds. per se or in the form of pharmaceutically acceptable salts. The compds. I and their salts can be employed as ingredients in pharmaceutical compns., optionally in combination with other antivirals, immunomodulators, antibiotics or vaccines.

IT 865300-19-2P

RN

RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of hydroxynaphthyridinediones as HIV integrase inhibitors) 865300-19-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo-, sodium salt (1:1) (CA INDEX NAME)

Na

IT 865298-58-4P 865298-70-0P 865298-71-1P

865300-11-4P 865300-12-5P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of hydroxynaphthyridinediones as HIV integrase inhibitors)

RN 865298-58-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(4-fluorophenyl)methyl]- 2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865298-70-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-N-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865298-71-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-N-(2,2,2-trifluoroethyl)- (CA INDEX NAME)

RN 865300-11-4 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 5-bromo-2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 865300-12-5 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy- (CA INDEX NAME)

865298-59-5P 865298-60-8P 865298-61-9P ΙT 865298-62-0P 865298-63-1P 865298-64-2P 865298-65-3P 865298-66-4P 865298-67-5P 865298-68-6P 865298-69-7P 865298-72-2P 865298-73-3P 865298-74-4P 865298-75-5P 865298-76-6P 865298-77-7P 865298-78-8P 865298-79-9P 865298-80-2P 865298-81-3P 865298-82-4P 865298-83-5P 865298-84-6P 865298-85-7P 865298-86-8P 865298-87-9P 865298-88-0P 865298-89-1P 865298-90-4P 865298-91-5P 865298-92-6P 865298-93-7P 865298-94-8P 865298-95-9P 865298-96-0P 865298-97-1P 865298-98-2P 865298-99-3P 865299-00-9P 865299-01-0P 865299-02-1P 865299-03-2P 865299-04-3P 865299-05-4P 865299-06-5P 865299-07-6P 865299-08-7P 865299-09-8P 865299-10-1P 865299-11-2P 865299-12-3P 865299-13-4P 865299-14-5P 865299-15-6P 865299-16-7P 865299-17-8P

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865299-22-5P 865299-24-7P 865299-25-8P
865299-26-9P 865299-27-0P 865299-28-1P
865299-29-2P 865299-30-5P 865299-31-6P
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865300-01-2P 865300-02-3P 865300-03-4P
865300-04-5P 865300-05-6P 865300-06-7P
865300-07-8P 865300-08-9P 865300-09-0P
865300-10-3P 865300-13-6P 865300-14-7P
865300-15-8P 865300-16-9P 865300-17-0P
865300-18-1P 865300-98-7P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)
   (preparation of hydroxynaphthyridinediones as HIV integrase inhibitors)
865298-59-5 CAPLUS
2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-
hexahydro-4-hydroxy-N, N-dimethyl-3, 5-dioxo- (CA INDEX NAME)
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RN

CN

RN 865298-60-8 CAPLUS
CN 2,6-Naphthyridine-1-carboxamide, N-cyclobutyl-6-[(4-fluorophenyl)methyl]2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo- (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ &$$

RN 865298-61-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclopropyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-(CA INDEX NAME)

RN 865298-62-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865298-63-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-64-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-(CA INDEX NAME)

RN 865298-65-3 CAPLUS

CN Methanesulfonamide, N-[6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl- (CA INDEX NAME)

RN 865298-66-4 CAPLUS

CN Acetamide, N-[6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl- (CA INDEX NAME)

RN 865298-67-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-68-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclobuty1-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-69-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclopropyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-72-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-N-[2-(methylsulfonyl)ethyl]-3,5-dioxo- (CA INDEX NAME)

RN 865298-73-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N,6-bis[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

$$\begin{array}{c|c} F \\ CH_2 \\ NH \\ C = O \\ Me \\ O \end{array}$$

RN 865298-74-4 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(1-piperidinylcarbonyl)- (CA INDEX NAME)

RN 865298-75-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-(2,2-dimethylpropyl)-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

$$\begin{array}{c|c} O \\ C-NH-CH_2-CMe_3 \\ \hline Me \\ O \\ OH \\ O \end{array}$$

RN 865298-76-6 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(4-thiomorpholinylcarbonyl)- (CA INDEX NAME)

RN 865298-77-7 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(1-piperazinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c} H \\ N \\ C = O \\ C \\ O \\ O \\ O \\ O \\ \end{array}$$

RN 865298-78-8 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(1-piperazinylcarbonyl)-, 2,2,2-trifluoroacetate (1:1) (CA INDEX NAME)

CM 1

CRN 865298-77-7 CMF C20 H21 F N4 O4

$$\begin{array}{c} & & \\$$

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 865298-79-9 CAPLUS

CN 1-Piperazinesulfonamide, 4-[[6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-2,6-naphthyridin-1-yl]carbonyl]-N,N-dimethyl-(CA INDEX NAME)

$$O = S - NMe_{2}$$

$$O = S - NMe_{2}$$

$$N$$

$$C = O$$

$$NH$$

$$O = NH$$

$$O = O$$

$$O = O$$

RN 865298-80-2 CAPLUS

CN 1-Piperazineacetamide, 4-[[6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-2,6-naphthyridin-1-yl]carbonyl]-N,N-dimethyl-α-oxo- (CA INDEX NAME)

RN 865298-81-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-82-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-, sodium salt (1:1) (CA INDEX NAME)

Na

RN 865298-83-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-84-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(2-methylpropyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-85-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-2-(2-methylpropyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-86-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N,N-diethyl-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-87-9 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-[(4-methyl-1-piperazinyl)carbonyl]- (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{N} \\ \text{C} \\ \text{O} \\ \text$$

RN 865298-88-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-[(4-methyl-1-piperazinyl)carbonyl]-, 2,2,2-trifluoroacetate (1:1) (CA INDEX NAME)

CM 1

CRN 865298-87-9

CMF C22 H24 C1 F N4 O4

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 865298-89-1 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(4-thiomorpholinylcarbonyl)- (CA INDEX NAME)

RN 865298-90-4 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(1-piperidinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\$$

RN 865298-91-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N-(cyclopropylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-92-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N-cyclopropyl-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-93-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N-ethyl-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-94-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-N-(1-methylethyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-95-9 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-[(4,4-difluoro-1-piperidinyl)carbonyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-(CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & &$$

RN 865298-96-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(4-morpholinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & \\ & & & \\ &$$

RN 865298-97-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-98-2 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-[(4-cyclopropyl-1-piperazinyl)carbonyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-(CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\$$

RN 865298-99-3 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-[(4-cyclopropyl-1-piperazinyl)carbonyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-,2,2,2-trifluoroacetate (1:1) (CA INDEX NAME)

CM 1

CRN 865298-98-2 CMF C24 H26 C1 F N4 O4

$$\begin{array}{c|c} & & & \\ & & & \\$$

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 865299-00-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N,N-diethyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-01-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-[2-(dimethylamino)ethyl]-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-02-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-N-(1-methyl-4-piperidinyl)-3,5-dioxo-(CA INDEX NAME)

RN 865299-03-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N,6-bis[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-3,5-dioxo-(CA INDEX NAME)

$$\begin{array}{c} F \\ CH2 \\ N-Me \\ C=0 \\ O \\ O \\ O \\ O \\ O \end{array}$$

RN 865299-04-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N,N-diethyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo- (CA INDEX NAME)

RN 865299-05-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-N-(2-methylpropyl)-3,5-dioxo- (CA INDEX NAME)

RN 865299-06-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-ethyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-07-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-3,5-dioxo-N-propyl- (CA INDEX NAME)

RN 865299-08-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-N-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865299-09-8 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(1-pyrrolidinylcarbonyl)- (CA INDEX NAME)

RN 865299-10-1 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(4-morpholinylcarbonyl)- (CA INDEX NAME)

RN 865299-11-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-6-[[3-(trifluoromethyl)phenyl]methyl]- (CA INDEX NAME)

RN 865299-12-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[[4-fluoro-3-(trifluoromethyl)phenyl]methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-13-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-(1,3-benzodioxol-4-ylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-14-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-(1,3-benzodioxol-5-ylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-15-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-6-[(2-methoxyphenyl)methyl]-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-16-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-6-[(3-methoxyphenyl)methyl]-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-17-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-6-[(3-methylphenyl)methyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-18-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,4-dimethylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-19-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2,3-dichlorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-20-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2,4-difluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-22-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-6-[[3-(trifluoromethoxy)phenyl]methyl]- (CA INDEX NAME)

RN 865299-24-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-25-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluoro-3-methylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-26-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2-bromo-3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-27-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-6-[(2-methylphenyl)methyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-28-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-29-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-6-[(4-methylphenyl)methyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-30-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-chlorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-31-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-6-[(4-methoxyphenyl)methyl]-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-32-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,5-dichlorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-33-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,4-dichlorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-34-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,5-difluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-35-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,5-dimethoxyphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-36-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-methylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-37-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-fluoro-4-methylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-38-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2,5-difluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-39-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-2-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-40-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-chloro-2-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-41-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(5-chloro-2-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-42-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2-fluoro-3-methylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-43-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(5-fluoro-2-methylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-44-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,5-dimethylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-45-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-6-[(4-hydroxyphenyl)methyl]-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-46-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2,3-dihydro-1,4-benzodioxin-6-yl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-47-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluoro-3-methoxyphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-48-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-methoxyphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-49-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-chloro-3-methylphenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-50-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3,4-difluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-51-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(2-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-52-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chlorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-53-2 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2,3,4,6-tetrahydro-8-hydroxy-2-[(4-methoxyphenyl)methyl]-6-methyl-5-(1-pyrrolidinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ \text{MeO} & & & \\ & & & \\ \text{CH}_2 & & & \\ & & & \\ \text{O} & & \\ \text{OH} & & \\ \end{array}$$

RN 865299-54-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,3,5,6,7,8-hexahydro-4-hydroxy-6-[(4-methoxyphenyl)methyl]-N,2-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-55-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-(1,3-benzodioxol-5-ylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865299-56-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(4-methylphenyl)-3,5-dioxo-(CA INDEX NAME)

RN 865299-57-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-2-phenyl-(CA INDEX NAME)

RN 865299-58-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluoropheny1)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-2-(3-thienyl)-(CA INDEX NAME)

RN 865299-59-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-2-(3-pyridinyl)-(CA INDEX NAME)

RN 865299-60-1 CAPLUS

CN Benzoic acid, 4-[6-[(3-chloro-4-fluorophenyl)methyl]-1[(dimethylamino)carbonyl]-5,6,7,8-tetrahydro-4-hydroxy-3,5-dioxo-2,6naphthyridin-2(3H)-yl]-, methyl ester (CA INDEX NAME)

RN 865299-61-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2-[4-(aminocarbonyl)phenyl]-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

$$H_2N-C$$
 $C-NMe_2$
 C
 N
 CH_2
 F

RN 865299-62-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[(methylamino)carbonyl]phenyl]-3,5-dioxo-(CA INDEX NAME)

RN 865299-63-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-[4-[(ethylamino)carbonyl]phenyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-64-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[[(1-methylethyl)amino]carbonyl]phenyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-65-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-[4-[(dimethylamino)carbonyl]phenyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-66-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-[4-[(diethylamino)carbonyl]phenyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-67-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-[3-[(dimethylamino)carbonyl]phenyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-68-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(4-nitrophenyl)-3,5-dioxo-(CA INDEX NAME)

RN 865299-69-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2-[4-(acetylamino)phenyl]-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-70-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2-[4-(acetylmethylamino)phenyl]-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-71-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[methyl(2,2,2-trifluoroacetyl)amino]phenyl]-3,5-dioxo-(CA INDEX NAME)

$$F_3C-C-N \qquad \qquad C-NMe_2 \qquad C1 \\ N \qquad \qquad N-CH_2 \qquad \qquad F$$

RN 865299-72-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[methyl[(methylamino)carbonyl]amino]phenyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-73-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-[4-[(dimethylamino)carbonyl]methylamino]phenyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-74-7 CAPLUS

CN Carbamic acid, [4-[6-[(3-chloro-4-fluorophenyl)methyl]-1[(dimethylamino)carbonyl]-5,6,7,8-tetrahydro-4-hydroxy-3,5-dioxo-2,6naphthyridin-2(3H)-yl]phenyl]-, methyl ester (9CI) (CA INDEX NAME)

RN 865299-75-8 CAPLUS

CN Carbamic acid, [4-[6-[(3-chloro-4-fluorophenyl)methyl]-1[(dimethylamino)carbonyl]-5,6,7,8-tetrahydro-4-hydroxy-3,5-dioxo-2,6naphthyridin-2(3H)-yl]phenyl]methyl-, methyl ester (9CI) (CA INDEX NAME)

RN 865299-76-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[(methylsulfonyl)amino]phenyl]-3,5-dioxo-(CA INDEX NAME)

RN 865299-77-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[methyl(methylsulfonyl)amino]phenyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-78-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2-[4-(aminosulfonyl)phenyl]-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-79-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-[(methylamino)sulfonyl]phenyl]-3,5-dioxo-(CA INDEX NAME)

RN 865299-80-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-(4-morpholinylsulfonyl)phenyl]-3,5-dioxo- (CA INDEX NAME)

RN 865299-81-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[4-(methylsulfonyl)phenyl]-3,5-dioxo-(CA INDEX NAME)

RN 865299-82-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(3-cyanophenyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-83-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(4-cyanophenyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-84-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2-(4-acetylphenyl)-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-85-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(cyanomethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-86-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(1-cyanoethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-87-2 CAPLUS

CN 2,6-Naphthyridine-2(3H)-acetamide, 6-[(3-chloro-4-fluorophenyl)methyl]-1[(dimethylamino)carbonyl]-5,6,7,8-tetrahydro-4-hydroxy-3,5-dioxo- (CA INDEX NAME)

RN 865299-88-3 CAPLUS

CN 2,6-Naphthyridine-2(3H)-acetamide, 6-[(3-chloro-4-fluorophenyl)methyl]-1[(dimethylamino)carbonyl]-5,6,7,8-tetrahydro-4-hydroxy-α-methyl-3,5dioxo- (CA INDEX NAME)

RN 865299-89-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(cyclopropylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-90-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(cyclobutylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-91-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-(cyclohexylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-92-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-(2-methoxyethyl)-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-93-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-2-(2,2,2-trifluoroethyl)- (CA INDEX NAME)

RN 865299-94-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-2-(phenylmethyl)-(CA INDEX NAME)

RN 865299-95-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865299-96-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2,6-bis[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-97-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-2-[2-(1-pyrrolidinyl)ethyl]- (CA INDEX NAME)

RN 865299-98-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-[2-(4-morpholinyl)ethyl]-3,5-dioxo-(CA INDEX NAME)

$$\begin{array}{c|c} O \\ C-NMe_2 \\ O \\ N-CH_2-CH_2-N \\ OH \\ O \\ \end{array}$$

RN 865299-99-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 2-(2-aminoethyl)-6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo- (CA INDEX NAME)

$$\begin{array}{c|c} O \\ C-NMe_2 \\ O \\ \hline \\ OH \\ O \end{array} \quad \begin{array}{c} C1 \\ F \\ \hline \\ CH_2 \\ \hline \\ \end{array}$$

RN 865300-00-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865300-01-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(2-methylpropyl)-3,5-dioxo- (CA INDEX NAME)

RN 865300-02-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(5-chloro-4-fluoro-2-iodophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865300-03-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(5-chloro-4-fluoro-2-iodophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(2-methylpropyl)-3,5-dioxo- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & \\ i-\text{Bu} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ &$$

RN 865300-04-5 CAPLUS

CN Methanesulfonamide, N-[6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl-(CA INDEX NAME)

RN 865300-05-6 CAPLUS

CN Ethanesulfonamide, N-[6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl-(CA INDEX NAME)

RN 865300-06-7 CAPLUS

CN Methanesulfonamide, N-[6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-(2-methylpropyl)-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl- (CA INDEX NAME)

RN 865300-07-8 CAPLUS

CN Methanesulfonamide, N-[6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-(2-methylpropyl)-3,5-dioxo-2,6-naphthyridin-1-yl]-N-ethyl- (CA INDEX NAME)

RN 865300-08-9 CAPLUS

CN Urea, N-[6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-2,6-naphthyridin-1-yl]-N,N',N'-trimethyl- (CA INDEX NAME)

RN 865300-09-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(5-methyl-1,3,4-oxadiazol-2-yl)- (CA INDEX NAME)

RN 865300-10-3 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 5-bromo-2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy- (CA INDEX NAME)

RN 865300-13-6 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 865300-14-7 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-ethyl-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 865300-15-8 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-cyclopropyl-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 865300-16-9 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(3-pyridinyl)- (CA INDEX NAME)

RN 865300-17-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(4-pyridinyl)- (CA INDEX NAME)

RN 865300-18-1 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-(2-furanyl)-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 865300-98-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6-tetrahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

IT 865300-88-5

RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of hydroxynaphthyridinediones as HIV integrase inhibitors)

RN 865300-88-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 2,3,5,6,7,8-hexahydro-4-hydroxy-6-[(4-methoxyphenyl)methyl]-2-methyl-3,5-dioxo-, methyl ester (CA INDEX NAME)

IT 865300-36-3P 865300-53-4P 865300-54-5P 865300-62-5P 865300-73-8P 865300-81-8P

865300-86-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of hydroxynaphthyridinediones as HIV integrase inhibitors)

RN 865300-36-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(4-fluorophenyl)methyl]2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, ethyl ester, sodium salt (1:1)
(CA INDEX NAME)

Na

RN 865300-53-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865300-54-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865300-62-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865300-73-8 CAPLUS

CN Carbamic acid, [2-[6-[(3-chloro-4-fluorophenyl)methyl]-1[(dimethylamino)carbonyl]-5,6,7,8-tetrahydro-4-hydroxy-3,5-dioxo-2,6naphthyridin-2(3H)-yl]ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

RN 865300-81-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-(CA INDEX NAME)

RN 865300-86-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, ethyl ester (CA INDEX NAME)

6

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2005:1026949 CAPLUS

DOCUMENT NUMBER: 143:326343

TITLE: Preparation of hydroxynaphthyridinediones as HIV

integrase inhibitors

INVENTOR(S): Han, Wei; Egbertson, Melissa; Wai, John S.; Zhuang,

Linghang; Ruzek, Rowena D.; Perlow, Debra S.;

Obligado, Vanessa E.

PATENT ASSIGNEE(S): Merck & Co., Inc., USA SOURCE: PCT Int. Appl., 83 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PA	PATENT NO.					KIND DATE		APPLICATION NO.					DATE					
WC	WO 2005087767					20050922		WO 2005-US7730					20050309					
	W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,	
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	ΚZ,	LC,	
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,	
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	
		SY,	ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW
	RW:	BW,	GH,	GM,	ΚE,	LS,	MW,	MZ,	NΑ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	
		ΑZ,	BY,	KG,	KΖ,	MD,	RU,	ТJ,	TM,	ΑT,	ΒE,	BG,	CH,	CY,	CZ,	DE,	DK,	
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IS,	ΙΤ,	LT,	LU,	MC,	NL,	PL,	PT,	
		RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	G₩,	ML,	
		MR,	NE,	SN,	TD,	ΤG												
BF	BR 2005008495				A	20070731			BR 2005-8495					20050309				
IN	IN 2006CN03154				Α	20070608			IN 2006-CN3154				20060831					
MX	MX 2006PA10252				Α	20061030			MX 2006-PA10252			20060908						
PRIORIT	RIORITY APPLN. INFO.:									US 2004-551625P				P 20040309				
										US 2	004-	6331	34P	:	P 2	0041	203	
										WO 2	005-	US77	30		W 2	0050	309	

OTHER SOURCE(S): MARPAT 143:326343

GΙ

The title compds. I [X1 = H, Br, Cl, F, OMe; X2 = H, Br, Cl, F, OMe, AΒ alkyl, etc.; R4 = CO2H, C(O)O(alkyl), CONH2, etc.; R5 = H, alkyl, cycloalkyl, etc.] which are inhibitors of HIV integrase and inhibitors of HIV replication, and therefore are useful in the prevention and treatment of infection by HIV and in the prevention, delay in the onset, and treatment of AIDS, were prepared and formulated. E.g., a multi-step synthesis of II, starting from piperidin-2-one and 4-fluorobenzyl bromide, was given. The representative compds. I (including II) were tested in the integrase assay and were found to have IC50's less than about 1 μM . The compds. I can be employed against HIV infection and AIDS as compds. per se or in the form of pharmaceutically acceptable salts. The compds. I and their salts can be employed as ingredients in pharmaceutical compns., optionally in combination with other antivirals, immunomodulators, antibiotics or vaccines.

ΙT 865300-19-2P

> RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of hydroxynaphthyridinediones as HIV integrase inhibitors)

865300-19-2 CAPLUS RN

2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-CN 2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo-, sodium salt (1:1) (CA INDEX NAME)

Na

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865298-70-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-N-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

IT 865298-59-5P 865298-60-8P 865298-61-9P 865298-62-0P 865298-63-1P 865298-64-2P 865298-65-3P 865298-66-4P 865298-67-5P 865298-68-6P 865298-69-7P 865298-74-4P 865298-76-6P 865298-77-7P 865298-78-8P 865298-81-3P 865298-82-4P 865298-83-5P

865298-84-6P 865298-85-7P 865298-86-8P 865298-87-9P 865298-88-0P 865298-89-1P 865298-90-4P 865298-91-5P 865298-92-6P 865298-93-7P 865298-94-8P 865298-95-9P 865298-96-0P 865298-97-1P 865299-00-9P 865299-04-3P 865299-05-4P 865299-06-5P 865299-07-6P 865299-08-7P 865299-09-8P 865299-10-1P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of hydroxynaphthyridinediones as HIV integrase inhibitors) RN 865298-59-5 CAPLUS 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-CN hexahydro-4-hydroxy-N, N-dimethyl-3, 5-dioxo- (CA INDEX NAME)

RN 865298-60-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclobutyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-(CA INDEX NAME)

RN 865298-61-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclopropyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo- (CA INDEX NAME)

RN 865298-62-0 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865298-63-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-64-2 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo- (CA INDEX NAME)

RN 865298-65-3 CAPLUS

CN Methanesulfonamide, N-[6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl- (CA INDEX NAME)

RN 865298-66-4 CAPLUS

CN Acetamide, N-[6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-2,6-naphthyridin-1-yl]-N-methyl- (CA INDEX NAME)

RN 865298-67-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-68-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclobuty1-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-69-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-cyclopropyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-74-4 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(1-piperidinylcarbonyl)- (CA INDEX NAME)

RN 865298-76-6 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(4-thiomorpholinylcarbonyl)- (CA INDEX NAME)

RN 865298-77-7 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(1-piperazinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c} H \\ N \\ C = O \\ \\ CH_2 - N \\ O \end{array}$$

RN 865298-78-8 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(1-piperazinylcarbonyl)-, 2,2,2-trifluoroacetate (1:1) (CA INDEX NAME)

CM 1

CRN 865298-77-7 CMF C20 H21 F N4 O4

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 865298-81-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-82-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N,2-trimethyl-3,5-dioxo-, sodium salt (1:1) (CA INDEX NAME)

Na

RN 865298-83-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(1-methylethyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-84-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-2-(2-methylpropyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-85-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-2-(2-methylpropyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-86-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluoropheny1)methyl]-N,N-diethyl-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo- (CA INDEX NAME)

RN 865298-87-9 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-[(4-methyl-1-piperazinyl)carbonyl]- (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \\ \text{N} \\ \\ \text{C} \\ \text{O} \\ \text{OH} \\ \end{array}$$

RN 865298-88-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-[(4-methyl-1-piperazinyl)carbonyl]-, 2,2,2-trifluoroacetate (1:1) (CA INDEX NAME)

CM 1

$$\begin{array}{c|c} & \text{Me} \\ & \\ & \\ \text{N} \\ & \\ \text{C} \\ & \\ \text{O} \\ & \text{OH} \\ \end{array}$$

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 865298-89-1 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(4-thiomorpholinylcarbonyl)- (CA INDEX NAME)

RN 865298-90-4 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(1-piperidinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c|c} C1 & C = 0 \\ \hline F & N \\ \hline CH_2 - N & O \\ \hline O & OH \\ \end{array}$$

RN 865298-91-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N-(cyclopropylmethyl)-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-92-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N-cyclopropyl-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865298-93-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-N-ethyl-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-3,5-dioxo-(CA INDEX

NAME)

RN 865298-94-8 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluoropheny1)methy1]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethy1-N-(1-methylethyl)-3,5-dioxo-(CA INDEX NAME)

RN 865298-95-9 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-5-[(4,4-difluoro-1-piperidinyl)carbonyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-(CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & &$$

RN 865298-96-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chloro-4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl-5-(4-morpholinylcarbonyl)- (CA INDEX NAME)

$$\begin{array}{c} C1 & C = O \\ N & Me \\ CH_2 - N & O \\ O & OH \end{array}$$

RN 865298-97-1 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,2-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-00-9 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N,N-diethyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-04-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N,N-diethyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-(CA INDEX NAME)

RN 865299-05-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-N-(2-methylpropyl)-3,5-dioxo- (CA INDEX NAME)

RN 865299-06-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, N-ethyl-6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-3,5-dioxo-(CA INDEX NAME)

RN 865299-07-6 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-3,5-dioxo-N-propyl- (CA INDEX NAME)

RN 865299-08-7 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N-methyl-N-(1-methylethyl)-3,5-dioxo- (CA INDEX NAME)

RN 865299-09-8 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(1-pyrrolidinylcarbonyl)- (CA INDEX NAME)

RN 865299-10-1 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(4-fluorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-5-(4-morpholinylcarbonyl)- (CA INDEX NAME)

IT 865300-36-3P 865300-53-4P 865300-54-5P

865300-62-5P 865300-86-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of hydroxynaphthyridinediones as HIV integrase inhibitors)

RN 865300-36-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(4-fluorophenyl)methyl]-

2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, ethyl ester, sodium salt (1:1) (CA INDEX NAME)

Na

RN 865300-53-4 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865300-54-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-

2,3,5,6,7,8-hexahydro-4-hydroxy-2-methyl-3,5-dioxo-, methyl ester (CA INDEX NAME)

RN 865300-62-5 CAPLUS

CN 2,6-Naphthyridine-1-carboxamide, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-N,N-dimethyl-3,5-dioxo-(CA INDEX NAME)

RN 865300-86-3 CAPLUS

CN 2,6-Naphthyridine-1-carboxylic acid, 6-[(3-chloro-4-fluorophenyl)methyl]-2,3,5,6,7,8-hexahydro-4-hydroxy-3,5-dioxo-, ethyl ester (CA INDEX NAME)

REFERENCE COUNT:

6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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ACCESSION NUMBER: 2005:158668 CAPLUS

DOCUMENT NUMBER: 142:261561

TITLE: Preparation of pyrido[1,2-a]pyrazine-1,8-dione

derivatives as HIV integrase inhibitors

INVENTOR(S): Miyazaki, Susumu; Katoh, Susumu; Adachi, Kaoru; Isoshima, Hirotaka; Kobayashi, Satoru; Matsuzaki,

Yuji; Watanabe, Wataru; Yamataka, Kazunobu; Kiyonari,

Shinichi; Wamaki, Shuichi

PATENT ASSIGNEE(S): Japan Tobacco Inc., Japan SOURCE: PCT Int. Appl., 355 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.					KIND		DATE			APPLICATION NO.				DATE				
WO	2005016927			A1	_	20050224			WO 2004-JP11869				20040812					
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ORITY APPLN. INFO.:									JP 2	003-	2931	17		A 2	0030	813		
										JP 2	004-	1348	96		A 2	0040	428	
								JP 2005-513209				A3 20040812						
										WO 2	004-	JP11	869	,	W 2	0040	812	
										US 2	004-	9582	25		B1 2	0041	005	

OTHER SOURCE(S): MARPAT 142:261561

AB The title compds. I [wherein R1 = (un)substituted alkyl, alkenyl, alkynyl, etc.; X = (un)substituted CH2, N=CH, or CH=N; Y1-Y2-Y3 = (un)substituted C=CH-NH, N-CH=N, N-CH=CH, C=N-NH, N-N=CH, etc.; R2 = H, alkyl, arylalkyl, or (un)substituted SO2H] or pharmaceutically acceptable salts thereof are

prepared as anti-HIV agents. For example, the compound II \bullet HCl was prepared in a multi-step synthesis. II \bullet HCl inhibited HIV integrase with IC50 of <0.01 μ M. Formulations containing I as an active ingredient were also described.

IT 845719-70-2P 845719-85-9P 845719-86-0P

845720-07-2P 845720-08-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of pyrido[1,2-a]pyrazine-1,8-dione derivs. as HIV integrase inhibitors)

RN 845719-70-2 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3-chlorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 845719-85-9 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[(3,4-dichlorophenyl)methyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

RN 845719-86-0 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2-[3-(2-chloro-6-fluorophenyl)propyl]-2,3,4,6-tetrahydro-8-hydroxy-6-methyl- (CA INDEX NAME)

Me
$$_{N}$$
 $_{OH}$ $_{O}$ $_{OH}$ $_{O}$ $_{F}$

RN 845720-07-2 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2,3,4,6-tetrahydro-8-hydroxy-6-methyl-2-[[3-(trifluoromethyl)phenyl]methyl]- (CA INDEX NAME)

RN 845720-08-3 CAPLUS

CN 2,6-Naphthyridine-1,7-dione, 2,3,4,6-tetrahydro-8-hydroxy-2-[(3-methoxyphenyl)methyl]-6-methyl- (CA INDEX NAME)

REFERENCE COUNT:

10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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